

## CLAIMS

1. A method of treatment of a patient undergoing opioid analgesic therapy which comprises minimising or mitigating the side effects of the opioid by the administration of a therapeutically effective amount of devazepide.

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2. A method of treatment of a patient requiring analgesia which comprises the administration of a therapeutically effective amount of an opioid analgesic whilst minimising the side effects of the opioid by the separate, simultaneous or sequential administration of a therapeutically effective amount of devazepide.

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3. A method according to claim 1 characterised in that the opioid is selected from the group morphine, or a salt thereof such as the sulphate, chloride or hydrochloride, or the other 1,4-hydroxymorphinan opioid analgesics such as naloxone, meperidine, butorphanol or pentazocine, or morphine-6-glucuronide, codeine, dihydrocodeine, diamorphine, dextropropoxyphene, pethidine, fentanyl, alfentanil, alphaprodine, buprenorphine, dextromoramide, diphenoxylate, dipipanone, heroin (diacetylmorphine), hydrocodone (dihydrocodeinone), hydromorphone (dihydromorphinone), levorphanol, meptazinol, methadone, metopon (methyldihydromorphinone), nalbuphine, oxycodone, 15 buprenorphine, dextromoramide, diphenoxylate, dipipanone, heroin (diacetylmorphine), hydrocodone (dihydrocodeinone), hydromorphone (dihydromorphinone), levorphanol, meptazinol, methadone, metopon (methyldihydromorphinone), nalbuphine, oxycodone, 20 phenazocine, remifentanil, tramadol, or a salt of any of these.

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4. A method according to claim 2 characterised in that the opioid is selected from the group morphine, or a salt thereof such as the sulphate, chloride or hydrochloride, or the other 1,4-hydroxymorphinan opioid analgesics such as naloxone, meperidine, butorphanol or pentazocine, or morphine-6-glucuronide, codeine, dihydrocodeine, diamorphine, dextropropoxyphene, pethidine, fentanyl, alfentanil, alphaprodine, buprenorphine, dextromoramide, diphenoxylate, dipipanone, heroin (diacetylmorphine), hydrocodone (dihydrocodeinone), hydromorphone (dihydromorphinone), levorphanol, meptazinol, methadone, metopon (methyldihydromorphinone), nalbuphine, oxycodone, 30 phenazocine, remifentanil, tramadol, or a salt of any of these.

(dihydrohydroxycodeinone), oxymorphone (dihydrohydroxymorphinone), phenadoxone, phenazocine, remifentanil, tramadol, or a salt of any of these

5. A method according to claim 3 characterised in that the opioid is selected from the group hydromorphone, oxycodone, morphine and fentanyl.

6. A method according to claim 4 characterised in that the opioid is selected from the group hydromorphone, oxycodone, morphine and fentanyl.

10 7. A method according to claim 5 characterised in that the opioid is selected from the group morphine and morphine sulphate.

8. A method according to claim 6 characterised in that the opioid is selected from the group morphine and morphine sulphate.

15 9. A method according to claim 1 characterised in that the method of delivery of the devazepide and/or the opioid is selected from the group, administration intravenously, orally, intrathecally, intranasally, intrarectally, intramuscularly/subcutaneously, by inhalation and by transdermal patch.

20 10. A method according to claim 2 characterised in that the method of delivery of the devazepide and/or the opioid is selected from the group, administration intravenously, orally, intrathecally, intranasally, intrarectally, intramuscularly/subcutaneously, by inhalation and by transdermal patch.

25 11. A method according to claim 9 characterised in that the devazepide is administered intravenously or orally.

30 12. A method according to claim 10 characterised in that the devazepide is administered intravenously or orally.

13. A method according to claim 11 characterised in that the devazepide is administered orally.

5 14. A method according to claim 12 characterised in that the devazepide is administered orally.

10 15. A method according to claim 9 characterised in that the opioid is administered intravenously and the devazepide is administered intravenously.

16. A method according to claim 10 characterised in that the opioid is administered intravenously and the devazepide is administered intravenously.

15 17. A method according to claim 9 characterised in that the opioid is administered orally and the devazepide is administered orally.

18. A method according to claim 10 characterised in that the opioid is administered orally and the devazepide is administered orally.

20 19. A method according to claim 9 characterised in that the opioid is administered by intravenous administration or oral administration.

20 21. A method according to claim 10 characterised in that the opioid is administered by intravenous administration or oral administration.

25 22. A method according to claim 1 characterised in that the daily dosage of devazepide is up to 0.7 mg/kg/day.

30 23. A method according to claim 2 characterised in that the daily dosage of devazepide is up to 0.7 mg/kg/day.

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23. A method according to claims 21 characterised in that the daily dosage of devazepide is from 25  $\mu$ g/kg/day to 0.7 mg/kg/day.

5 24. A method according to claims 22 characterised in that the daily dosage of devazepide is from 25  $\mu$ g/kg/day to 0.7 mg/kg/day.

10 25. A method according to claim 23 characterised in that the daily dosage of devazepide is from 50  $\mu$ g/kg/day to 0.5 mg/kg/day.

26. A method according to claim 24 characterised in that the daily dosage of devazepide is from 50  $\mu$ g/kg/day to 0.5 mg/kg/day.

15 27. A method according to claim 25 characterised in that the devazepide is administered orally and the daily dosage of devazepide is from 0.07 mg/kg/day to 0.29 mg/kg/day.

20 28. A method according to claim 26 characterised in that the devazepide is administered orally and the daily dosage of devazepide is from 0.07 mg/kg/day to 0.29 mg/kg/day.

25 29. A method according to either of claims 25 characterised in that the devazepide is administered intravenously administration the dosage of devazepide is from 50  $\mu$ g/kg/day to 0.5 mg/kg/day.

30. A method according to either of claims 26 characterised in that the devazepide is administered intravenously administration the dosage of devazepide is from 50  $\mu$ g/kg/day to 0.5 mg/kg/day.

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31. A method according to claim 1 characterised in that the daily dosage of the opioid is from 5 to 2000mg daily.

32. A method according to claim 2 characterised in that the daily dosage of the opioid is from 5 to 2000mg daily.

33. A method according to claim 31 characterised in that the daily dosage of the opioid is from 5 to 100mg daily.

10 34. A method according to claim 32 characterised in that the daily dosage of the opioid is from 5 to 100mg daily.

35. A method according to claim 1 characterised in that the side effect which is inhibited, mitigated or minimised is selected from the group, constipation, dizziness, 15 tiredness/fatigue and vomiting.

36. A method according to claim 2 characterised in that the side effect which is inhibited, mitigated or minimised is selected from the group, constipation, dizziness, tiredness/fatigue and vomiting.

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37. A method according to claim 1 characterised in that the devazepide used is the S enantiomer wherein the level of R enantiomer is not greater than 1.5% w/w.

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38. A method according to claim 2 characterised in that the devazepide used is the S enantiomer wherein the level of R enantiomer is not greater than 1.5% w/w.

39. The use of devazepide in the manufacture of a medicament which inhibits or mitigates the undesirable side effects of administration of a therapeutically effective amount of an opioid analgesic.

40. The use according to claim 39 characterised in that the devazepide used is the S enantiomer wherein the level of R enantiomer is not greater than 1.5% w/w.

5 41. The use of devazepide in the manufacture of a medicament for use in the method  
of either of claim 1.

42. The use of devazepide in the manufacture of a medicament for use in the method of either of claim 2.

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